

FORM PTO-1500 (REV. 9-2001)		U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE		ATTORNEY'S DOCKET NUMBER	
TRANSMITTAL LETTER TO THE UNITED STATES DESIGNATED/ELECTED OFFICE (DO/EO/US) CONCERNING A FILING UNDER 35 U.S.C. 371				34156	
				U.S. APPLICATION NO. (if known, see 37 CFR 1.5) <b>09/979576</b>	
INTERNATIONAL APPLICATION NO. PCT/GB00/01937		INTERNATIONAL FILING DATE 19 May 2000		PRIORITY DATE CLAIMED 20 May 1999	
TITLE OF INVENTION CHROMATOGRAPHY COLUMNS					
APPLICANT(S) FOR DO/EO/US Martin John Hofmann					
Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:					
<p>1. <input checked="" type="checkbox"/> This is a <b>FIRST</b> submission of items concerning a filing under 35 U.S.C. 371.</p> <p>2. <input type="checkbox"/> This is a <b>SECOND</b> or <b>SUBSEQUENT</b> submission of items concerning a filing under 35 U.S.C. 371.</p> <p>3. <input type="checkbox"/> This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (21) indicated below.</p> <p>4. <input checked="" type="checkbox"/> The US has been elected by the expiration of 19 months from the priority date (Article 31).</p> <p>4. <input checked="" type="checkbox"/> A copy of the International Application as filed (35 U.S.C. 371(c)(2))</p> <p>a. <input type="checkbox"/> is attached hereto (required only if not communicated by the International Bureau).</p> <p>b. <input checked="" type="checkbox"/> has been communicated by the International Bureau.</p> <p>c. <input type="checkbox"/> is not required, as the application was filed in the United States Receiving Office (RO/US).</p> <p>5. <input type="checkbox"/> An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)).</p> <p>a. <input checked="" type="checkbox"/> is attached hereto.</p> <p>b. <input type="checkbox"/> has been previously submitted under 35 U.S.C. 154(d)(4).</p> <p>6. <input checked="" type="checkbox"/> Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3))</p> <p>a. <input type="checkbox"/> are attached hereto (required only if not communicated by the International Bureau).</p> <p>b. <input type="checkbox"/> have been communicated by the International Bureau.</p> <p>c. <input type="checkbox"/> have not been made; however, the time limit for making such amendments has NOT expired.</p> <p>d. <input checked="" type="checkbox"/> have not been made and will not be made.</p> <p>7. <input type="checkbox"/> An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371(c)(3)).</p> <p>8. <input type="checkbox"/> An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)).</p> <p>10. <input type="checkbox"/> An English language translation of the annexes of the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)).</p> <p><b>Items 11 to 20 below concern document(s) or information included:</b></p> <p>11. <input checked="" type="checkbox"/> An Information Disclosure Statement under 37 CFR 1.97 and 1.98.</p> <p>12. <input type="checkbox"/> An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.</p> <p>13. <input checked="" type="checkbox"/> A FIRST preliminary amendment.</p> <p>14. <input type="checkbox"/> A SECOND or SUBSEQUENT preliminary amendment.</p> <p>15. <input type="checkbox"/> A substitute specification.</p> <p>16. <input type="checkbox"/> A change of power of attorney and/or address letter.</p> <p>17. <input type="checkbox"/> A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 35 U.S.C. 1.821 - 1.825.</p> <p>18. <input type="checkbox"/> A second copy of the published international application under 35 U.S.C. 154(d)(4).</p> <p>19. <input type="checkbox"/> A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4).</p> <p>20. <input checked="" type="checkbox"/> Other items or information:          --PCT Request; --PCT/RO/105;          --PCT/IB/301, 304 &amp; 308;          --PCT Demand; --Int'l. Prel. Exam. Report          --Publication No. WO 00/72002</p>					

Express Mail No. EL609746754US

I hereby certify that this correspondence is being deposited with the United States Postal Service "Express Mail Post Office to Addressee" service under 37 CFR 1.10, and is addressed to: Assistant Commissioner for Patents, Washington D.C. 20231 on the date indicated below.

Debra A. Peterlin

Name of Person Mailing Correspondence

11/19/2001 *Debra A. Peterlin*  
Date Signature of Person  
Mailing Correspondence

U.S. PATENT AND TRADE OFFICE <b>09-979576</b>		INTERNATIONAL APPLICATION NO. <b>PCT/GB00/01937</b>		ATTORNEY'S DOCKET NUMBER <b>34156</b>	
--	--	--	--	--	--

21. <input checked="" type="checkbox"/> The following fees are submitted: <b>BASIC NATIONAL FEE (37 CFR 1.492 (a) (1) - (5)):</b> Neither international preliminary examination fee (37 CFR 1.482) nor international search fee (37 CFR 1.445(a)(2)) paid to USPTO and International Search Report not prepared by the EPO or JPO ..... <b>\$1040.00</b> International preliminary examination fee (37 CFR 1.482) not paid to USPTO but International Search Report prepared by the EPO or JPO ..... <b>\$890.00</b> International preliminary examination fee (37 CFR 1.482) not paid to USPTO but international search fee (37 CFR 1.445(a)(2)) paid to USPTO ..... <b>\$740.00</b> International preliminary examination fee (37 CFR 1.482) paid to USPTO but all claims did not satisfy provisions of PCT Article 33(1)-(4) ..... <b>\$710.00</b> International preliminary examination fee (37 CFR 1.482) paid to USPTO and all claims satisfied provisions of PCT Article 33(1)-(4) ..... <b>\$100.00</b> <b>ENTER APPROPRIATE BASIC FEE AMOUNT =</b>			<b>CALCULATIONS PTO USE ONLY</b>		
			<b>890.00</b>		
			<b>\$890.00</b>		
Surcharge of <b>\$130.00</b> for furnishing the oath or declaration later than <input type="checkbox"/> 20 <input checked="" type="checkbox"/> 30 months from the earliest claimed priority date (37 CFR 1.492(e)).			<b>\$130.00</b>		
CLAIMS	NUMBER FILED	NUMBER EXTRA	RATE	\$	
Total claims	- 20 =		x <b>\$18.00</b>	\$	
Independent claims	- 3 =		x <b>\$84.00</b>	\$	
MULTIPLE DEPENDENT CLAIM(S) (if applicable)			+ <b>\$280.00</b>	\$ <b>\$280.00</b>	
<b>TOTAL OF ABOVE CALCULATIONS =</b>			<b>\$1,300.00</b>		
<input type="checkbox"/> Applicant claims small entity status. See 37 CFR 1.27. The fees indicated above are reduced by 1/2.			+		
<b>SUBTOTAL =</b>			<b>\$1,300.00</b>		
Processing fee of <b>\$130.00</b> for furnishing the English translation later than <input type="checkbox"/> 20 <input type="checkbox"/> 30 months from the earliest claimed priority date (37 CFR 1.492(f)).			\$		
<b>TOTAL NATIONAL FEE =</b>			<b>\$1,300.00</b>		
Fee for recording the enclosed assignment (37 CFR 1.21(h)). The assignment must be accompanied by an appropriate cover sheet (37 CFR 3.28, 3.31). <b>\$40.00</b> per property +			\$		
<b>TOTAL FEES ENCLOSED =</b>			<b>\$1,300.00</b>		
			Amount to be refunded:		\$
			charged:		\$

a. ☒ A check in the amount of \$ 1,300.00 to cover the above fees is enclosed.

b. ☐ Please charge my Deposit Account No. \_\_\_\_\_ in the amount of \$ \_\_\_\_\_ to cover the above fees. A duplicate copy of this sheet is enclosed.


c. ☒ The Commissioner is hereby authorized to charge any additional fees which may be required, or credit any overpayment to Deposit Account No. 16-0820. A duplicate copy of this sheet is enclosed.

d. ☐ Fees are to be charged to a credit card. **WARNING:** Information on this form may become public. Credit card information should not be included on this form. Provide credit card information and authorization on PTO-2038.

**NOTE:** Where an appropriate time limit under 37 CFR 1.494 or 1.495 has not been met, a petition to revive 37 CFR 1.137 (a) or (b)) must be filed and granted to restore the application to pending status.

SEND ALL CORRESPONDENCE TO:  
 Pearne & Gordon, LLP  
 526 Superior Avenue East  
 Suite 1200  
 Cleveland, Ohio 44114-1484

  
 SIGNATURE  
 Howard G. Shimola  
 NAME  
 26232  
 REGISTRATION NUMBER

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant: Martin John Hofmann  
Title: CHROMATOGRAPHY COLUMNS  
Docket No.: 34156

PRELIMINARY AMENDMENT

Assistant Commissioner for Patents  
Washington, D.C. 20231

Sir:

Prior to the examination of the above-identified patent application, it is requested that the following amendments be made.

IN THE CLAIMS:

Please amend claims 6, 8 and 9 to read as follows:

1 6. (amended) A chromatography column according to  
2 claim 1, in which an outwardly-directed sealing portion  
3 at or adjacent the front end of the plunger which makes a  
4 seal directly against the column wall, or which mounts a  
5 deformable seal element for making such a seal, is joined  
6 to the permeable filter portion via a one-piece integral  
7 structure.

1 8. (amended) A chromatography column according to  
2 claim 1, in which the plunger further comprises an outer  
3 plunger wall spaced outwardly from said tubular stem  
4 defining the internal flow conduit, the outer plunger  
5 wall and tubular stem being integrally bonded to one

09979576-021502

6 another at the front end of the plunger so as to seal off  
7 an internal space of the plunger, around said tubular  
8 stem, at the front end.

1 9. (amended) A chromatography column according to  
2 claim 1, in which one end of the column tube has a full-  
3 diameter opening receiving the plunger and the other end  
4 is a closed end, converging to a union for an external  
5 fluid flow conduit and having a fixed permeable filter  
6 element across the column tube adjacent the closed end.

**REMARKS**

The foregoing amendments correct multiple claim dependency for purposes of calculating the claim fee.

Attached hereto are pages entitled "Version With Markings to Show Changes Made".

If there are any fees required by this amendment not covered by an enclosed check, or if no check is enclosed, please charge the same to Deposit Account No. 16-0820, Order No. 34156.

Respectfully submitted,

By: 

Howard G. Shimola, Reg. No. 26232

526 Superior Avenue East  
Suite 1200  
Cleveland, Ohio 44114-1484  
(216) 579-1700

November 19, 2001

"VERSION WITH MARKINGS TO SHOW CHANGES MADE"

IN THE CLAIMS:

Claims 6, 8 and 9 have been amended in the following manner:

09979576-021502  
1           6. (amended) A chromatography column according to  
2 [any one of the preceding claims] claim 1, in which an  
3 outwardly-directed sealing portion at or adjacent the  
4 front end of the plunger which makes a seal directly  
5 against the column wall, or which mounts a deformable  
6 seal element for making such a seal, is joined to the  
7 permeable filter portion via a one-piece integral  
8 structure.

1           8. (amended) A chromatography column according to  
2 [any one of the preceding claims] claim 1, in which the  
3 plunger further comprises an outer plunger wall spaced  
4 outwardly from said tubular stem defining the internal  
5 flow conduit, the outer plunger wall and tubular stem  
6 being integrally bonded to one another at the front end  
7 of the plunger so as to seal off an internal space of the  
8 plunger, around said tubular stem, at the front end.

1           9. (amended) A chromatography column according to  
2 [any one of the preceding claims] claim 1, in which one  
3 end of the column tube has a full-diameter opening

4 receiving the plunger and the other end is a closed end,  
5 converging to a union for an external fluid flow conduit  
6 and having a fixed permeable filter element across the  
7 column tube adjacent the closed end.

09979576-021502

21PRTS

- 1 -

CHROMATOGRAPHY COLUMNSFIELD OF THE INVENTION

This invention has to do with chromatography columns and in particular laboratory columns, i.e. small columns such as are used for bench scale or research work.

BACKGROUND OF THE INVENTION

Generally speaking a column of the relevant kind (referred to herein as being "of the kind described") has a column tube and end filter arrangements which, in use, retain a bed of particulate chromatography medium in the column tube between them while allowing the passage of fluid for chromatography. At least one of the end filter arrangements is at the front end of a plunger which is axially slidable along inside the tubular column, makes a seal outwardly against the column tube and incorporates an internal flow conduit communicating along the plunger between a filter portion of the filter arrangement and a rear part of the plunger outside the column tube.

PRIOR ART

Columns of the kind described are currently available at varying levels of cost and complexity.

The column tube is usually glass.

In sophisticated versions the plunger shaft has outer and inner stiff concentric plastic tubes. The inner tube may be a thick solid tube, with a narrow bore providing the flow conduit. Or it may be a wide-bored tube housing an inmost fine flexible tube which is the flow conduit. At the front end of this shaft a rubber sealing ring is trapped between end formations of the inner and outer stiff tubes, and the filter arrangement has a porous disc (e.g. a

5 sinter or mesh) clamped over a divergent flow distributing  
end surface of the inner tube, at the outlet of the flow  
conduit. The outer stiff tube runs slidably through a  
plastics end unit screwed onto the end of the column tube.  
10 The end unit has a mechanism which may be switched between  
a free sliding engagement with the plunger's outer plastics  
tube (for large axial movements) and a screw-threaded  
engagement (for fine adjustments). An interned screw  
engagement is also provided between the inner and outer  
tubes so that after advancing the plunger to the desired  
position - i.e. contacting the end of the bed of medium -  
the rubber sealing ring can be squeezed out into sealing  
contact with the column tube wall by axially compressing it  
between the end formations of the inner and outer tubes.  
15 Internal plunger seals are also provided to prevent leakage  
from between the internal conduit and the filter element  
into the interior of the plunger construction, or to the  
outside of the shaft. These columns give good results but  
can be very expensive, and are complicated to use and  
20 maintain.

Laboratory columns of another, simpler kind use short  
removable plugs at both ends of the column. The plugs are  
solid polymeric units having an outer O-ring seal to seal  
against the glass column wall, a narrow central bore for  
25 the fluid flow and a flat annular recess at the inner face  
which receives the porous filter disc. The plug has a very  
short travel into the column tube, so the media bed must be  
carefully filled to the correct depth. The necessarily  
tight-fitting seal makes it hard work to push the plug into  
30 the column end. The outside of the glass tube end is



threaded to take a clamping nut which holds the plug in once installed. These columns are simpler and cheaper than known plunger columns of the kind described, but less versatile.

5 SUMMARY OF THE INVENTION

A first proposal herein is that the plunger of a column of the kind described comprises a tubular stem of glass or other formable material, preferably transparent, which defines in one piece the internal fluid flow conduit. The permeable filter element is integrally bonded to the front end of this tubular stem across the internal fluid flow conduit.

In particular the filter element is preferably bonded to the plunger stem by being integrally fused therewith, e.g. by heat-fusing. The materials of the filter element and stem can be selected for compatibility in this respect, e.g. both may be of glass or a suitable thermoplastics material. A preferred version has a glass stem fused to a sintered glass filter element.

20 Preferably the tubular stem defining the internal fluid flow conduit extends as a one-piece integral whole rearwardly to a rear connection union at the rear of the plunger, i.e. a threaded union, spigot or ferrule. Preferably this union has a joint boundary which is exterior of the plunger stem. Like the integral bonding of the filter element at the front end, this one-piece construction provides a simple means for preventing leakage within the plunger which is a significant difficulty with prior art constructions.

30 A further proposal, preferably combined with the

above, is that a one-piece integral construction joins the permeable filter element and an outwardly-directed sealing portion at or adjacent the front end of the plunger which makes a seal directly against the column tube wall, or which mounts a deformable seal element for making such a seal.

In particular the filter element is preferably bonded to the plunger's outer wall by being integrally fused therewith, e.g by heat-fusing. Again, use of glass or thermoplastics material for the filter element and plunger outer wall is advantageous in this respect; a sintered glass filter element may be fused to an outer glass wall of the plunger.

A further independent proposal herein, again preferably combined with the above, is that the plunger comprises a tubular stem of glass or other formable material which defines, preferably in one piece, the internal fluid flow conduit, and also an outer plunger wall spaced outwardly from the tubular stem, the outer plunger wall and tubular stem being integrally bonded to one another at the front end of the plunger so as to seal off the internal space of the plunger at the front end. Preferably one or both of the stem and plunger outer wall is/are integrally bonded to the filter element, as in the first and/or second aspects set out above. Again the use of fused glass or other thermoplastics is advantageous. Prevention of leakage to within the plunger structure can therefore be prevented reliably without additional internal sealing components. By combination with the other aspects above, the entire internal fluid flow path from a rear

connection union to the permeable filter element can be sealed vis á vis the plunger interior without requiring seal components between discrete mechanical parts. The use of glass or other suitable thermoplastics enables such a construction to be made easily e.g using conventional plastics-forming or glass-blowing techniques. Furthermore the use of a transparent material enables the user to observe the flow of material within the plunger.

An outwardly-directed sealing portion at or adjacent the front end of the plunger may have a plunger wall surface which is shaped e.g by machining or moulding, most preferably a machined glass surface, to fit and seal directly against the column tube wall. The use of ground glass surfaces to make fluid-tight seals is as such well-known in the laboratory context. The present proposal can exploit this in a new way, enabling a special advantage in combination with the other aspects as described above and the transparency available with glass material. Machined glass surfaces are not highly transparent but become so when in wetted contact with another glass surface. The plunger exterior may thus have a cylindrical sealing portion making a fitting seal against the column tube wall interior. This sealing portion may be axially elongate, so as to align the plunger axially in the column and avoid the need for column tube end units as were required in the prior art. The same advantage may be achieved by other plunger constructions providing an axially-elongate fitting engagement inside the column tube, e.g axially-extending fins or the like behind the sealing portion which may itself then be shorter.

The plunger may be made with a deformable e.g resilient sealing element such as a sealing ring, preferably an O-ring seal, fitting around the plunger's outer wall to make a sealing arrangement against the column tube wall.

The internal fluid flow conduit preferably has an elongate portion of relatively narrow cross section (e.g less than 1 or 2mm diameter) and a divergent (larger cross-section) distribution portion immediately adjacent to the permeable filter element. Where the internal tubular conduit is provided in a discrete tubular stem within an outer plunger wall, the tubular stem is preferably flared at its lower end to provide this divergence. Again this is easy to do with glass or thermoplastics tubular stems.

Particularly in smaller size columns, the plunger may be essentially a solid rod whose outer surface opposes the column wall, with a narrow central bore for the fluid flow conduit. In this case the front end of the rod may be shaped to form a divergent zone around the front opening of the flow bore.

The plunger can resemble that of a syringe. It may have a head at its rear (outward) end adapted for manual pushing. Preferably a rear connection union for the internal fluid flow conduit emerges transversely from the plunger, below such a head.

The column tube wall may be double-ended as in known constructions, and plungers as described above may be deployed at both ends. More preferably however we propose that one end of the column tube wall has a full-diameter opening receiving the mentioned plunger and the other end

is a closed end, converging to a union for an external fluid flow conduit, with a fixed permeable filter element across the column tube adjacent the closed end. The fixed permeable element may be permanently installed, e.g a glass sinter disc fused into a glass column tube wall. Or, a mount may be provided for fixed mounting of an exchangeable filter element.

The volume of the column is typically not more than 100ml.

#### BRIEF DESCRIPTION OF THE DRAWINGS

Embodiments of the invention are described with reference to the accompanying drawings, in which

FIG 1 is an axial cross-sectional view of a laboratory chromatography column embodying the invention, and

FIGS 2-5 illustrate some possible variants of plunger construction.

#### DETAILED DESCRIPTION OF PREFERRED EMBODIMENTS

With reference to Fig 1, an all-glass laboratory chromatography column consists of a column tube 1 in which a plunger 2 fits slidably

The column tube 1 in this embodiment is the tube of a commercially-available syringe. Its volume is for example 50ml. At its upper open end it has a conventional flange 12 with a flat 17 at one side to prevent rolling. The column has the usual uniform cylindrical wall 11, which can conveniently be mounted upright in a laboratory clamp 6. The lower end of the column has an integral convergence 13 to a nozzle 15, in which a metal spigot 16 is incorporated to form an outlet union connectable, e.g by a conventional ferrule, to an outlet pipe if desired.

A sintered glass filter disc 5 is fixedly mounted across the lower end of the tube 1 at the transition to the convergent portion 13. The tube wall at this portion 14 has been heat deformed and fused with the periphery of the glass sintered disc 5 to form a permanent joint.

The glass column wall 11 is transparent in the usual way.

The plunger 2 is also an all-glass construction, made from standard laboratory glass using standard glass-forming techniques. Its three main components are a central tubular stem 32 defining a flow conduit 3, an outer tubular housing wall 22 surrounding the stem and having a top head 21, and a sintered glass permeable filter disc 4 mounted in the front end of the stem.

The tubular outer wall 22 has a lower portion extending approximately half the length of the plunger, nearly to the front end, which has a machined glass surface 23 making a sliding sealing fit against the inner surface of the column tube wall 11. By this means the plunger 2 defines with the column tube 1 a sealed internal bed space 7 between the upper and lower filter elements 4,5. No resilient seal element is required, although one may be used if desired adjacent to the front end. The elongate close-fitting glass surface 23 also maintains axial alignment of the plunger 2 in the column 1. Above a shoulder 24 at the top of the machined surface 23 the plunger 22 narrows, so that it can be slid as far down as desired inside the column tube 1. A chromatography medium bed of any desired length can therefore be established in the space 7. A laboratory clamp 6' suffices to hold the

plunger down in contact on top of the medium bed.

5 The internal tubular stem 32 of the plunger can be made from stock glass tubing e.g of 1mm bore. Using standard glass forming techniques its lower end is flared outwardly to form a divergent section 33 and its periphery is fused onto the upper surface of the glass sinter disc 4. A fused joint is also established between the mouth 25 of the outer glass wall 22 and the periphery of the sinter disc 4, and onto the edge of the divergent portion 33 of the central stem. By exploiting glass' intrinsic propensity for forming and fusing by simple hot manipulations, this provides a plunger end construction which is fully internally sealed (i.e preventing leaks into the space between the stem 32 and outer wall 22) and externally sealed against the column wall 11, without requiring any discrete sealing elements. It is also fully transparent, so that the user can observe the flow of sample through the conduit 3 and down onto the sinter disc 4, verifying the absence of trapped air in the divergent space 35 above the disc 4 - this latter is important, and difficult to check in conventional opaque constructions.

10 It is in generally desirable to minimise the volume of the distinction space 35 and not difficult to do so in thermoformable material. The same applies to the lower convergence 13 of the column tube.

15 The surface groove around the outer <sup>a</sup>well behind the joint to the disc will provide a useful seating for a discrete deformable seal element, either additionally to the glass-glass seal illustrated or, in a alternative embodiment, or the sole external seal.

At its upper end the central tubular stem 32 is bent sideways and emerges through the outer tubular wall 22, below the enlarged head 21. The end of the tube 32 is bonded into this opening by a thermally-fused joint 38, completing the sealing of the fluid flow path in the plunger *vis á vis* the exterior. The upper opening of the tube mouth 37 has a connection union formed with an internal glass thread 36. Alternatively, the tube may project laterally beyond the column wall 22 as a nozzle to house a metal spigot of the kind used at the bottom of the column 1, or some other suitable union may be used.

It will be noted that the head 21 is clear of the union opening 27 so that the plunger can be manipulated without interfering with the fluid connections.

The skilled reader will appreciate that the various integrally-fused bonds created in this embodiment, exploiting the properties of glass, individually provide advantages over prior art constructions. Useful embodiments of our proposals can therefore be made with only some of these integral bonds, and/or using different materials. The double-tube construction shown in Fig 1 may be too elaborate for very small columns, and Fig 2 shows the end of a plunger for a small column, in which a single hollow glass rod provides both the exterior surface 23 and the internal conduit 3 of the plunger. In this very simple structure the divergence at the front end of the conduit is dispensed with and the sinter disc 4 is fused or otherwise bonded directly onto the front face of the tubular rod 39.

Fig 3 shows a similar solid-rod construction, modified by the machining of a divergent chamber 35 immediately



upstream of the permeable disc 4, and the recessing of the disc 4 into the front face of the rod 39.

In the Fig 2 and Fig 3 constructions the outer surface 23 of the rod effects the seal against the tube wall. Fig 4 shows how additionally or alternatively a deformable seal element such as a rubber O-ring 223 or a PTFE ring may be fitted around the front end of the plunger exterior.

In all of the above embodiments a full cylindrical contact of the plunger in the column tube assures the axial alignment of the two main components. However this is not essential, and particularly in plastics constructions material may be saved by using a construction as in Fig 5. Here a one-piece moulded plastics entity forming the plunger has a central tubular portion 332 defining the internal conduit 3, surrounded by axially-extending fins 222 which provide the necessary axial alignment against the column tube wall, and a solid end unit 250 providing the necessary seal against the tube wall, e.g via PTFE ring 223, and the surrounding fused or bonded join to the permeable filter element 4.

CLAIMS

1. A chromatography column having a column tube and end filter arrangements which, in use, retain a bed of particulate chromatography medium in the column tube between them while allowing the passage of fluid for chromatography;

at least one of the end filter arrangements being at the front end of a plunger which is axially slidable along inside the column tube, makes a seal outwardly against the tube and incorporates an internal flow conduit communicating along the plunger between a permeable filter portion of the respective end filter arrangement and a rear part of the plunger outside the column tube;

the plunger comprising a tubular stem of glass or other formable material which defines in one piece said internal flow conduit, the permeable filter portion being integrally bonded to the front end of the tubular stem across the internal flow conduit.

2. A chromatography column according to claim 1 in which the filter portion is integrally fused to the plunger stem.

3. A chromatography column according to claim 1 or claim 2 in which both stem and filter portion are of glass or thermoplastics material.

4. A chromatography column according to claim 1 in which the tubular stem extends as a one-piece integral whole back to a rear connection union at the rear of the plunger.

5. A chromatography column according to claim 4 in which the rear connection union has a joint boundary at the exterior of the plunger stem.

6. A chromatography column according to any one of the preceding claims in which an outwardly-directed sealing portion at or adjacent the front end of the plunger which makes a seal directly against the column wall, or which mounts a deformable seal element for making such a seal, is joined to the permeable filter portion via a one-piece integral instruction.

7. A chromatography column according to claim 6 in which the permeable filter portion is bonded to the plunger's outer wall by being integrally fused therewith.

8. A chromatography column according to any one of the preceding claims in which the plunger further comprises an outer plunger wall spaced outwardly from said tubular stem defining the internal flow conduit, the outer plunger wall and tubular stem being integrally bonded to one another at the front end of the plunger so as to seal off an internal space of the plunger, around said tubular stem, at the front end.

9. A chromatography column according to any one of the preceding claims in which one end of the column tube has a full-diameter opening receiving the plunger and the other end is a closed end, converging to a union for an external fluid flow conduit and having a fixed permeable filter element across the column tube adjacent the closed end.

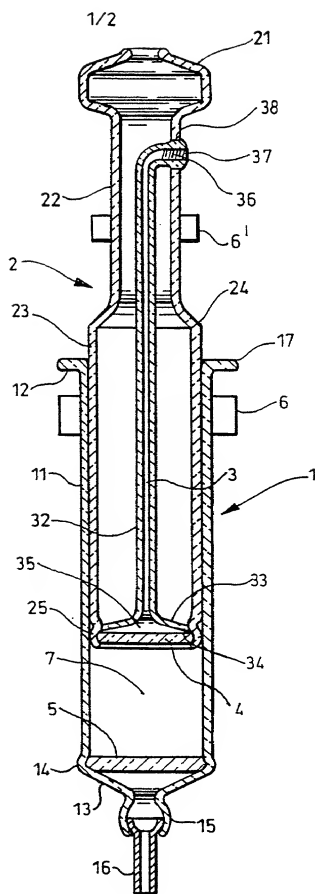


FIG.1.

2/2

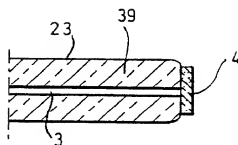


FIG. 2.

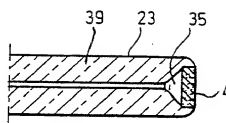


FIG. 3.

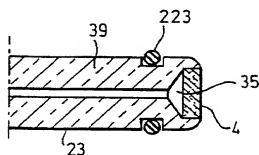


FIG. 4.

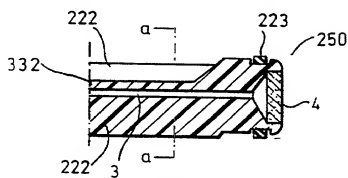


FIG. 5.

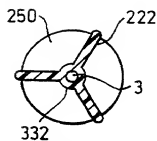


FIG. 5a.

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number

**DECLARATION FOR UTILITY OR  
DESIGN  
PATENT APPLICATION  
(37 CFR 1.63)**Declaration  
Submitted  
with Initial  
Filing

OR

Declaration  
Submitted after Initial  
Filing (surcharge  
(37 CFR 1.16 (e))  
required)

Attorney Docket Number 34156

First Named Inventor Martin John Hofmann

**COMPLETE IF KNOWN**

Application Number /

Filing Date

Art Unit

Examiner Name

As the below named inventor, I hereby declare that:

My residence, mailing address, and citizenship are as stated below next to my name.

I believe I am the original and first inventor of the subject matter which is claimed and for which a patent is sought on the invention entitled:

CHROMATOGRAPHY COLUMNS

(Title of the Invention)

the specification of which



is attached hereto

OR



was filed on (MM/DD/YYYY)

05/19/2000

as United States Application Number or PCT International

Application Number PCT/GB00/01937 and was amended on (MM/DD/YYYY) 11/19/2001 (if applicable).

I hereby state that I have reviewed and understand the contents of the above identified specification, including the claims, as amended by any amendment specifically referred to above.

I acknowledge the duty to disclose information which is material to patentability as defined in 37 CFR 1.56, including for continuation-in-part applications, material information which became available between the filing date of the prior application and the national or PCT international filing date of the continuation-in-part application.

I hereby claim foreign priority benefits under 35 U.S.C. 119(a)-(d) or (f), or 365(b) of any foreign application(s) for patent, inventor's or plant breeder's rights certificate(s), or 365(a) of any PCT international application which designated at least one country other than the United States of America, listed below and have also identified below, by checking the box, any foreign application for patent, inventor's or plant breeder's rights certificate(s), or any PCT international application having a filing date before that of the application on which priority is claimed.

Prior Foreign Application Number(s)	Country	Foreign Filing Date (MM/DD/YYYY)	Priority Not Claimed	Certified Copy Attached?	
				YES	NO
9911797.0	Great Britain	05/20/1999	<input type="checkbox"/>	<input type="checkbox"/>	<input checked="" type="checkbox"/>
PCT/GB00/01937	PCT	05/19/2000	<input type="checkbox"/>	<input type="checkbox"/>	<input checked="" type="checkbox"/>
			<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>


☐ Additional foreign application numbers are listed on a supplemental priority data sheet PTO/SB/02B attached hereto.

[Page 1 of 2]

Burden Hour Statement: This form is estimated to take 21 minutes to complete. Time will vary depending upon the needs of the individual case. Any comments on the amount of time you are required to complete this form should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, Washington, DC 20231. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO, Assistant Commissioner for Patents, Washington, DC 20231.

09770576.021502

## DECLARATION — Utility or Design Patent Application

Direct all correspondence to: <input checked="" type="checkbox"/> Customer Number or Bar Code Label				OR <input type="checkbox"/> Correspondence address below	
00116 PATENT TRADEMARK OFFICE					
Name					
Address					
City			State		ZIP
Country		Telephone		Fax	
<p>I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under 18 U.S.C. 1001 and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.</p>					
NAME OF SOLE OR FIRST INVENTOR: <input type="checkbox"/> A petition has been filed for this unsigned inventor					
Given Name (first and middle [if any]) <u>Martin John</u>			Family Name or Surname <u>Hofmann</u>		
Inventor's Signature <u>M. J. H.</u>			Date <u>16<sup>th</sup> Jan 2002</u>		
Residence: City <u>Stroud, Gloucestershire</u>		State	Great Britain		Citizenship <u>GBX</u>
Unit 4, Brimscombe Port Business Park, Brimscombe Hill, Brimscombe, Stroud, Mailing Address Gloucestershire GL5 2QN, Great Britain					
City		State	ZIP		Country
NAME OF SECOND INVENTOR: <input type="checkbox"/> A petition has been filed for this unsigned inventor					
Given Name (first and middle [if any])			Family Name or Surname		
Inventor's Signature			Date		
Residence: City		State	Country		Citizenship
Mailing Address					
City		State	ZIP		Country
<input type="checkbox"/> Additional inventors are being named on the ____ supplemental Additional Inventor(s) sheet(s) PTO/SB/02A attached hereto.					